

WHAT IS CLAIMED IS:

- 1        1. An isolated stem cell sustainable in culture under glycolytic conditions  
2 and which maintains the potential to differentiate.
- 1        2. The stem cell of claim 1 which is unipotent or pluripotent.
- 1        3. The stem cell of claim 1 which is an embryonic or somatic stem cell.  
1        4. The stem cell of claim 3 which is a pluripotent cell from a  
2 preimplantation embryo.
- 1        5. The stem cell of claim 1 which is a primordial germ cell.
- 1        6. The stem cell of claim 1 selected from the group consisting of  
2 hematopoietic, neuronal and mesenchymal stem cells.
- 1        7. An isolated stem cell which cell shows characteristic green staining  
2 with the mitochondrial marker JC-1.
- 1        8. An isolated stem cell which cell displays a low mitochondrial inner  
2 membrane potential based upon JC-1 green staining.
- 1        9. An isolated stem cell which cell displays a high mitochondrial inner  
2 membrane potential based upon JC-1 red staining.
- 1        10. A method of isolating a stem cell, comprising the steps of:
  - 2        (a) isolating a blastocyst;
  - 3        (b) identifying those cells which rely upon glycolysis for survival;
  - 4        (c) isolating a glycolytic cell from the inner cell mass of the blastocyst;
  - 5        and
  - 6        (d) culturing the isolated glycolytic cell to obtain an isolated stem cell.
- 1        11. The method of claim 10, wherein the cells are identified by staining  
2 with the mitochondrial marker JC-1.
- 1        12. The method of claim 10, further comprising maintaining the isolated  
2 cells on a fibroblast feeder layer to prevent differentiation.

- 1                   13. A chimeric animal produced from a cell of claims 1 or 9.
- 1                   14. A method of producing a chimeric animal comprising  
2                   (a) isolating a blastocyst;  
3                   (b) identifying those cells which rely upon glycolysis for survival;  
4                   (c) isolating the glycolytic cells from the inner cell mass of the blastocyst;  
5                   (d) transfecting a desired gene into the glycolytic cells;  
6                   (e) injecting the transfected cells into recipient blastocysts;  
7                   (f) implanting the transformed blastocysts into a host uterus; and  
8                   (g) nurturing the blastocysts to develop to term.
- 1                   15. A method of producing glycolytic-dependent cells, comprising the  
2 steps of:  
3                   (a) culturing cells under hypoxic conditions;  
4                   (b) identifying those cells which rely upon glycolysis for survival;  
5                   (c) isolating the glycolytic cells from the culture; and  
6                   (d) culturing the isolated glycolytic cells.
- 1                   16. A stem cell of claims 1 or 9 which is a mammalian stem cell.
- 1                   17. A chimeric mammal produced from a stem cell of claim 16.
- 1                   18. An isolated stem cell, wherein said stem cell can be identified by  
2 staining said cell with the fluorescent dye JC-1.
- 1                   19. The isolated stem cell of claim 18, wherein said cell is sensitive to  
2 inhibitors of multidrug resistance (MDR) targets.
- 1                   20. The isolated stem cell of claim 19, wherein said inhibitors are selected  
2 from the group consisting of verapamil, reserpine, and cyclosporine A.
- 1                   21. The isolated stem cell of claim 19, wherein the multidrug resistance  
2 (MDR) target is an MDR-like dye efflux pump.
- 1                   22. A method of identifying functionally distinct stem cells, comprising:  
2                   (a) staining the cells with the fluorescent dye JC-1;

3 (b) sorting the stained cells by fluorescence activated cell sorting  
4 (FACS);  
5 (c) analyzing said functionally distinct stem cells by comparing  
6 their sensitivity to inhibitors of multidrug resistance (MDR) targets; and  
7 (d) identifying a MDR-inhibitor sensitive JC-1 subpopulation of  
8 cells.

1                   23.     The MDR-inhibitor sensitive JC-1 subpopulation of claim 22, wherein  
2     said subpopulation has an increased differentiation permissiveness.

1                   24.     A method of switching embryonic stem cells between two  
2     subpopulations, comprising:

3 a) exposing a JC-1 green subpopulation to inhibitors of multidrug  
4 resistance genes; and  
5 b) overexpressing recombinant multidrug resistance genes in a JC-  
6 red subpopulation.

1                   25. The method of claim 24, wherein said inhibitors are selected from the  
2 group consisting of verapamil, reserpine and cyclosporine.

1                           26. A method of changing a cell's ability to differentiate by switching the  
2 subpopulations of claim 24.

1 27. An embryonic stem cell which is differentiated by the method of claim  
2 24.